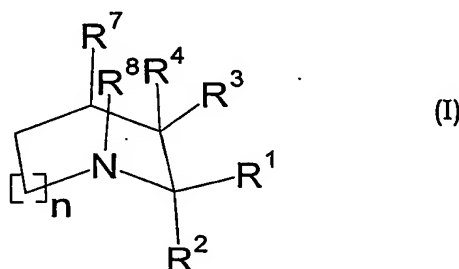


# Claims

1. Use of a compound capable of transferring wild type p53 from an inactive conformation thereof, which conformation is reactive to Pab 240 and not to Pab 1620,  
 5 into an active conformation capable of inducing apoptosis, which compound is selected from compounds having a structure according to the formula I



20 wherein

n is 0, 1 or 2;

R<sup>1</sup> and R<sup>2</sup> are the same or different and are selected from -H, -CH<sub>2</sub>-R<sup>5</sup>, -CH<sub>2</sub>-O-R<sup>5</sup>,

-CH<sub>2</sub>-S-R<sup>5</sup>, -CH<sub>2</sub>-NH-R<sup>5</sup>, -CO-O-R<sup>5</sup>, -CO-NH-R<sup>5</sup>, -CH<sub>2</sub>-NH-CO-R<sup>5</sup>,

25 -CH<sub>2</sub>-O-CO-R<sup>5</sup>, -CH<sub>2</sub>-NH-CO-NHR<sup>5</sup>, -CH<sub>2</sub>-NH-CO-OR<sup>5</sup>, -CH<sub>2</sub>-NH-CS-NHR<sup>5</sup> and -CH<sub>2</sub>-O-CO-NHR<sup>5</sup>; or R<sup>1</sup> and R<sup>2</sup> are together =CH<sub>2</sub>;

R<sup>3</sup> and R<sup>4</sup> are the same or different and are selected from -H, -OH, -SH, -NH<sub>2</sub>, -NHR<sup>5</sup> and -O-CO-C<sub>6</sub>H<sub>5</sub>; or R<sup>3</sup> and R<sup>4</sup> together are =O, =S, =NH or =NR<sup>5</sup>;

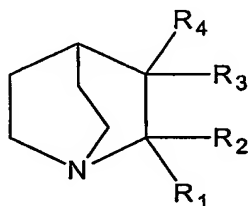
R<sup>5</sup> represents the same or different groups selected from H, substituted or  
 30 non-substituted C1 to C10 alkyl, C2 to C10 alkenyl, C2 to C10 alkynyl, substituted or non-substituted C3 to C12 cycloalkyl, substituted or non-substituted benzyl groups, substituted or non-substituted aryl or mono-, bi-, tricyclic unsubstituted or substituted heteroaromatic ring(s) with one or more heteroatoms and non-aromatic heterocycles wherein

35 the substituents of the substituted groups are selected from C1 to C10 alkyl, C2 to C10 alkenyl, C2 to C10 alkynyl, halogen, substituted or non-substituted aryl, substituted or non-substituted hetero-aromatic compounds, non-aromatic heterocycles, C1 to C10 alkyloxy, C1 to C10 alkylamino, C2 to C10 alkenylamino, C2 to C10 alkynylamino, COR<sup>6</sup>, CONR<sup>6</sup> and COOR<sup>6</sup>;

R<sup>6</sup> is selected from H, unsubstituted or substituted C1 to C10 alkyl, C2 to C10 alkenyl or alkynyl, benzyl, aryl, unsubstituted or substituted heteroaromatic rings with one or more hetero-atoms and non-aromatic heterocycles;

R<sup>7</sup> and R<sup>8</sup> together form a bridging CH<sub>2</sub>-CH<sub>2</sub> moiety; or R<sup>7</sup> and R<sup>8</sup> are both  
 5 hydrogen;  
 or a pharmaceutically acceptable salt or prodrug thereof,  
 for the preparation of a medicament for use in treating malignant melanoma  
 and/or a pathological condition involving undesired angiogenesis.

10 2. The use of claim 1, wherein the compound is selected from compounds having  
 the following formula (II)



(II)

15 wherein:

R<sub>1</sub> and R<sub>2</sub> are independently selected from hydrogen, hydroxymethyl, or  
 a methylene group linked to the nitrogen atom of an amine-substituted phenyl  
 group, to a nitrogen atom contained in the ring structure of a purine, 8-azapurine,  
 or benzimidazol residue, or R<sub>1</sub> and R<sub>2</sub> may together represent a double bonded me-  
 20 thylene group, and;

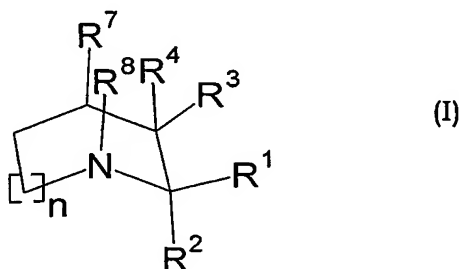
R<sub>3</sub> and R<sub>4</sub> are independently selected from hydrogen, hydroxyl, and  
 benzoyloxy, or R<sub>3</sub> and R<sub>4</sub> may together represent an oxygen atom being double  
 bonded, with the proviso that when either of R<sub>3</sub> and R<sub>4</sub> is a benzoyloxy group, both  
 R<sub>1</sub> and R<sub>2</sub> are hydrogen, or a pharmaceutically acceptable salt or prodrug thereof.

25 3. The use of claim 2, wherein the compound is selected from 2,2-  
 bis(hydroxymethyl)-1-azabicyclo[2.2.2]octan-3-one, 9-(azabicyclo[2.2.2]octan-3-  
 one)-6-chloro-9H-purine, 2-(hydroxymethyl)quinuclidine-3,3-diol, 2-(adenine-9-  
 methylene)-3-quinuclidinone, 2-methylene-3-quinuclidinone, 2-(-2-amino-3-chloro-  
 30 5-trifluoromethyl-1-methylaniline)-3-quinuclidinone, 2-(6-trifluoromethyl-4-  
 chlorobenzimidazole-1-methylene)-3-quinuclidinone, 2-(6-methoxypurine-9-

methylene)-3-quinuclidinone, 2-(8-azaadenine-9-methylene)-3-quinuclidinone, 1-azabicyclo [2.2.2]oct-3-yl benzoate, 2-(5,6-dimethyl-benzimidazole-1-methylene)-3-quinuclidinone, 2-(8-azaadenine-7-methylene)-3-quinuclidinone, 2-(7-methylene-1,3-dimethyluric acid)-3-quinuclidinone, or 2-(2,6-dichloro-9-methylenepurine)-3-quinuclidinone, or a pharmaceutically acceptable salt thereof.

4. The use of anyone of the claims 1-3 together with a pharmaceutically acceptable carrier, diluent and/or excipient.

5. A method of treating malignant melanoma and/or inhibiting undesired angiogenesis, comprising administering to a mammal in need thereof a pharmaceutically efficient amount of a compound selected from compounds having a structure according to the formula I



wherein

$n$  is 0, 1 or 2;

$R^1$  and  $R^2$  are the same or different and are selected from  $-H$ ,  $-CH_2-R^5$ ,  $-CH_2-O-R^5$ ,  $-CH_2-S-R^5$ ,  $-CH_2-NH-R^5$ ,  $-CO-O-R^5$ ,  $-CO-NH-R^5$ ,  $-CH_2-NH-CO-R^5$ ,  $-CH_2-O-CO-R^5$ ,  $-CH_2-NH-CO-NHR^5$ ,  $-CH_2-NH-CO-OR^5$ ,  $-CH_2-NH-CS-NHR^5$  and  $-CH_2-O-CO-NHR^5$ ; or  $R^1$  and  $R^2$  are together  $=CH_2$ ;

$R^3$  and  $R^4$  are the same or different and are selected from  $-H$ ,  $-OH$ ,  $-SH$ ,  $-NH_2$ ,  $-NHR^5$  and  $-O-CO-C_6H_5$ ; or  $R^3$  and  $R^4$  together are  $=O$ ,  $=S$ ,  $=NH$  or  $=NR^5$ ;

$R^5$  represents the same or different groups selected from  $H$ , substituted or non-substituted C1 to C10 alkyl, C2 to C10 alkenyl, C2 to C10 alkynyl, substituted or non-substituted C3 to C12 cycloalkyl, substituted or non-substituted benzyl groups, substituted or non-substituted aryl or mono-, bi-, tricyclic unsubstituted or substituted heteroaromatic ring(s) with one or more heteroatoms and non-aromatic heterocycles wherein

the substituents of the substituted groups are selected from C1 to C10 alkyl, C2 to C10 alkenyl, C2 to C10 alkynyl, halogen, substituted or non-substituted aryl, substituted or non-substituted hetero-aromatic compounds, non-aromatic heterocycles, C1 to C10 alkyloxy, C1 to C10 alkylamino, C2 to C10 alkenylamino, C2 to C10 alkynylamino, COR<sup>6</sup>, CONR<sup>6</sup> and COOR<sup>6</sup>;

R<sup>6</sup> is selected from H, unsubstituted or substituted C1 to C10 alkyl, C2 to C10 alkenyl or alkynyl, benzyl, aryl, unsubstituted or substituted heteroaromatic rings with one or more hetero-atoms and non-aromatic heterocycles;

R<sup>7</sup> and R<sup>8</sup> together form a bridging CH<sub>2</sub>-CH<sub>2</sub> moiety; or R<sup>7</sup> and R<sup>8</sup> are both hydrogen; or a pharmaceutically acceptable salt or prodrug thereof.

6. Method of testing compounds for the ability of transferring wild type p53 from an inactive conformation into an active conformation comprising the steps:

- A. Providing cells carrying wt p53, in which cells inactive wt p53 conformation is present;
- B. Exposing the cells *in vitro* to a substance to be tested; and
- C. Measuring the cellular inactive wt p53 conformation.

7. The method of claim 6, wherein instead of step C an alternative step C' is used comprising comparing the effect of the tested substance on the cells (carrying functional p53) in step B to the effect on cells or tissues with no or non-functional p53.

8. The method of claim 6 or 7, wherein integrin  $\alpha_v\beta_3$  is present in the cells.

9. The method of claim 6-8, wherein the Pab 240 is used for detecting wt p53 in its inactive conformation.

10. The method of any of the claims 6-9, wherein a compound of claim 1 is tested.

11. The method of any of the claims 6-10, wherein the cells in step B are exposed *in vivo* in an animal to the substance to be tested, and the animal subsequently sacrificed.